

CLAIMS

1. Purified polypeptide, comprising an amino acid sequence selected from the group consisting of:

- a) the sequence SEQ ID No. 2;
- b) the sequence SEQ ID No. 4;
- c) the sequence SEQ ID No. 6;
- d) the sequence SEQ ID No. 8;
- e) the sequence SEQ ID No. 10;
- f) the sequence SEQ ID No. 13;
- g) the sequence SEQ ID No. 15;
- h) the sequence SEQ ID No. 17;
- i) the sequence SEQ ID No. 19;

and j) any biologically active sequence derived from SEQ ID No. 2, SEQ ID No. 4, SEQ ID No. 6, SEQ ID No. 8, SEQ ID No. 10, SEQ ID No. 13, SEQ ID No. 15, SEQ ID No. 17 or SEQ ID No. 19.

2. Polypeptide according to Claim 1, characterized in that it comprises the amino acid sequence selected from the group consisting of SEQ ID No. 6, SEQ ID No. 13, SEQ ID No. 15, SEQ ID No. 17 and SEQ ID No. 19.

3. Polypeptide according to Claim 1, characterized in that it comprises the sequence lying between:

- residue 110 and residue 310 of SEQ ID No. 2 or 6;
- residue 60 and residue 260 of SEQ ID No. 8.

4. Polypeptide according to Claim 1, characterized in that it results from an alternative splicing of the messenger RNA of the corresponding gene.

5. Polypeptide according to any one of the preceding claims, characterized in that it is a recombinant polypeptide produced in the form of a fusion protein.

6. Isolated nucleic acid sequence coding for a polypeptide according to any one of the preceding claims.

7. Isolated nucleic acid sequence according to Claim 6, characterized in that it is selected from the group consisting of :

- a) the sequence SEQ ID No. 1;
- b) the sequence SEQ ID No. 3;
- c) the sequence SEQ ID No. 5;

- d) the sequence SEQ ID No. 7;
e) the sequence SEQ ID No. 9;
f) the sequence SEQ ID No. 11;
g) the sequence SEQ ID No. 12;
h) the sequence SEQ ID No. 14;
i) the sequence SEQ ID No. 16;
j) the sequence SEQ ID No. 18;
k) the nucleic acid sequences capable of hybridizing specifically with the sequence SEQ ID No. 1, SEQ ID No. 3, SEQ ID No. 5, SEQ ID No. 7, SEQ ID No. 9, SEQ ID No. 11, SEQ ID No. 12, SEQ ID No. 14, SEQ ID No. 16 or SEQ ID No. 18 or with the sequences complementary to them, or of hybridizing specifically with their proximal sequences;
and l) the sequences derived from the sequences a), b), c), d), e), f), g), h), i), j) or k) as a result of the degeneracy of the genetic code, mutation, deletion, insertion, and alternative splicing or an allelic variability.

8. Nucleotide sequence according to Claim 6, characterized in that it is a sequence selected from SEQ ID No. 5, SEQ ID No. 12, SEQ ID No. 14, SEQ ID No. 16 and SEQ ID No. 18 coding, respectively, for the polypeptide of sequences SEQ ID No. 6, SEQ ID No. 13, SEQ ID No. 15, SEQ ID No. 17 and SEQ ID No. 19.

9. Cloning and/or expression vector containing a nucleic acid sequence according to any one of Claims 6 to 8.

10. Vector according to Claim 9, characterized in that it is the plasmid pSE1.

11. Host cell transfected by a vector according to Claim 9 or 10.

12. Transfected host cell according to Claim 11, characterized in that it is *E. coli* MC 1061.

13. Nucleotide probe or nucleotide primer, characterized in that it hybridizes specifically with any one of the sequences according to Claims 6 to 8 or the sequences complementary to them or the corresponding messenger RNAs or the corresponding genes.

14. Probe or primer according to Claim 13, characterized in that it contains at least 16 nucleotides.

15. Probe or primer according to Claim 13, characterized in that it comprises the whole of the sequence of the gene coding for one of the polypeptides of Claim 1.

16. Nucleotide probe or primer selected from the group consisting of the following oligonucleotides or sequences complementary to them:

SEQ ID No. 20: GCG AGC TGC CCT CGG AG

SEQ ID No. 21: GGT TCT GCA GGT GAC TCA G

SEQ ID No. 22: GCC ATG CCT GTC TAC AAG

SEQ ID No. 23: ACC AGC TGG TTG ACG GAG

15 SEQ ID No. 24: GTC AAC CAG CTG GTG GGC CAG

SEQ ID No. 25: GTG GAT CTC GGC CTC C

SEQ ID No. 26: AGG CCG GCG TGG GGA AG

SEQ ID No. 27: CTT GGC GAT CTG GCA GTA G

SEQ ID No. 28: GCG GCC ACG ACC GTG AC

20 SEQ ID No. 29: GGC AGC TTG GGT CTC TGG

SEQ ID No. 30: CTG TAC GTC GGT GAC CCC

SEQ ID No. 31: TCA GTG GAT CTC GGC CTC

SEQ ID No. 32: AGG GGA CGC AGC GAA ACC

SEQ ID No. 33: CCA TCA GCT CCA GGC TCT C

25 SEQ ID No. 34: CCA GGA CAG GCG CAG ATG

SEQ ID No. 35: GAT GAG GTG GCT GGC TGG A

SEQ ID No. 36: TGG TCA GGT TCT GCA GGT G

SEQ ID No. 37: CAC CTA CTC CAG GGA TGC

SEQ ID No. 38: AGG AAA ATA GAA GCG TCA GTC

30 SEQ ID No. 39: CAG GCC CAC TTG CCT GCC

and SEQ ID No. 40: CTG TCC CCA AGC TGA TGA G

17. Use of a sequence according to any one of Claims 6 to 8, for the manufacture of oligonucleotide primers for sequencing reactions or specific amplification reactions according to the PCR technique or any variant of the latter.

18. Nucleotide primer pair, characterized in that it comprises the primers selected from the group consisting of the following sequences:

- a) sense primer: GCG AGC TGC CCT CGG AG (SEQ ID No. 20)
 antisense primer: GGT TCT GCA GGT GAC TCA G (SEQ ID No. 21)
- b) sense primer: GCC ATG CCT GTC TAC AAG (SEQ ID No. 22)
 antisense primer: ACC AGC TGG TTG ACG GAG (SEQ ID No. 23)
- c) sense primer: GTC AAC CAG CTG GTG GGC CAG (SEQ ID No. 24)
 antisense primer: GTG GAT CTC GGC CTC C (SEQ ID No. 25)
- d) sense primer: AGG CCG GCG TGG GGA AG (SEQ ID No. 26)
 antisense primer: CTT GGC GAT CTG GCA GTA G (SEQ ID No. 27)
- e) sense primer: GCG GCC ACG ACC GTG A (SEQ ID No. 28)
 antisense primer: GGC AGC TTG GGT CTC TGG (SEQ ID No. 29)
- f) sense primer: CTG TAC GTC GGT GAC CCC (SEQ ID No. 30)
 antisense primer: TCA GTG GAT CTC GGC CTC (SEQ ID No. 31)
- g) sense primer: AGG GGA CGC AGC GAA ACC (SEQ ID No. 32)
 antisense primer: GGC AGC TTG GGT CTC TGG (SEQ ID No. 29)
- h) sense primer: CCCCCCCCCCCCCCN (where N equals G, A or T)
 antisense primer: CCA TCA GCT CCA GGC TCT C (SEQ ID No. 33)
- i) sense primer: CCCCCCCCCCCCCCN (where N equals G, A or T)
 antisense primer: CCA GGA CAG GCG CAG ATG (SEQ ID No. 34)
- j) sense primer: CCCCCCCCCCCCCCN (where N equals G, A or T)
 antisense primer: CTT GGC GAT CTG GCA GTA G (SEQ ID No. 27)
- k) sense primer: CAC CTA CTC CAG GGA TGC (SEQ ID No. 37)
 antisense primer: AGG AAA ATA GAA GCG TCA GTC (SEQ ID No. 38)
- and l) sense primer: CAG GCC CAC TTG CCT GCC (SEQ ID No. 39)
 antisense primer: CTG TCC CCA AGC TGA TGA G (SEQ ID No. 40)
19. Use of a sequence according to any one of Claims 6 to 8, which is usable in gene therapy.
20. Use of a sequence according to any one of Claims 6 to 8, for the production of diagnostic

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nucleotide probes or primers, or of antisense sequences which are usable in gene therapy.

21. Use of nucleotide primers according to any one of Claims 6 to 8, for sequencing.

5 22. Use of a probe or primer according to any one of Claims 13 to 16, as an in vitro diagnostic tool for the detection, by hybridization experiments, of nucleic acid sequences coding for a polypeptide according to any one of Claims 1 to 4, in biological samples, or for
10 the demonstration of aberrant syntheses or of genetic abnormalities.

23. Method of in vitro diagnosis for the detection of aberrant syntheses or of genetic abnormalities in the nucleic acid sequences coding for a polypeptide
15 according to any one of Claims 1 to 4, characterized in that it comprises:

- the bringing of a nucleotide probe according to any one of Claims 13 to 16 into contact with a biological sample under conditions permitting the formation of a hybridization complex between the said probe and the abovementioned nucleotide sequence, where appropriate after a prior step of amplification of the abovementioned nucleotide sequence;
- 20 - the detection of the hybridization complex possibly formed;
- where appropriate, the sequencing of the nucleotide sequence forming the hybridization complex with the probe of the invention.

24. Use of a nucleic acid sequence according to any one of Claims 6 to 8, for the production of a recombinant polypeptide according to any one of Claims 1 to 5.

25. Method of production of a recombinant SR-p70
35 protein, characterized in that transfected cells according to Claim 10 or 11 are cultured under conditions permitting the expression of a recombinant polypeptide of sequence SEQ ID No. 2, SEQ ID No. 4, SEQ ID No. 6, SEQ ID No. 8, SEQ ID No. 10, SEQ ID No. 13,
40 SEQ ID No. 15, SEQ ID No. 17 or SEQ ID No. 19 or any

biologically active fragment or derivative, and in that the said recombinant polypeptide is recovered.

26. Mono- or polyclonal antibodies or their fragments, chimeric antibodies or immunoconjugates, characterized in that they are capable of specifically recognizing a polypeptide according to any one of Claims 1 to 4.

27. Use of the antibodies according to the preceding claim, for the purification or detection of a polypeptide according to any one of Claims 1 to 4 in a biological sample.

28. Method of in vitro diagnosis of pathologies correlated with an expression or an abnormal accumulation of SR-p70 proteins, in particular the phenomena of carcinogenesis, from a biological sample, characterized in that at least one antibody according to Claim 25 is brought into contact with the said biological sample under conditions permitting the possible formation of specific immunological complexes between an SR-p70 protein and the said antibody or antibodies, and in that the specific immunological complexes possibly formed are detected.

29. Kit for the in vitro diagnosis of an expression or an abnormal accumulation of SR-p70 proteins in a biological sample and/or for measuring the level of expression of these proteins in the said sample, comprising:

- at least one antibody according to Claim 25, optionally bound to a support,
- means of visualization of the formation of specific antigen-antibody complexes between an SR-p70 protein and the said antibody, and/or means of quantification of these complexes.

30. Method for the early diagnosis of tumour formation, characterized in that autoantibodies directed against an SR-p70 protein are demonstrated in a serum sample drawn from an individual, according to the steps that consist in bringing a serum sample drawn from an individual into contact with a polypeptide of the invention, optionally bound to a support, under

conditions permitting the formation of specific immunological complexes between the said polypeptide and the autoantibodies possibly present in the serum sample, and in that the specific immunological complexes possibly formed are detected.

31. Method of determination of an allelic variability, a mutation, a deletion, an insertion, a loss of heterozygosity or a genetic abnormality of the SR-p70 gene, characterized in that it utilizes at least one nucleotide sequence according to any one of Claims 6 to 8.

32. Method of determination of an allelic variability of the SR-p70 gene at position -30 and -20 relative to the initiation ATG of exon 2 which may be involved in pathologies, and characterized in that it comprises at least:

- a step during which exon 2 of the SR-p70 gene carrying the target sequence is amplified by PCR using a pair of oligonucleotide primers according to any one of Claims 6 to 8;
- a step during which the amplified products are treated with a restriction enzyme whose cleavage site corresponds to the allele sought;
- a step during which at least one of the products of the enzyme reaction is detected or assayed.

33. Pharmaceutical composition comprising as active principle a polypeptide according to any one of Claims 1 to 4.

34. Pharmaceutical composition according to the preceding claim, characterized in that it comprises a polypeptide according to Claim 2.

35. Pharmaceutical composition containing an inhibitor or an activator of SR-p70 activity.

36. Pharmaceutical composition containing a polypeptide derived from a polypeptide according to any one of Claims 1 to 5, characterized in that it is an inhibitor or an activator of SR-p70.